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CARRIERS OF THE KIF6 719ARG ALLELE APPEAR TO HAVE A HIGHER LP(A) CONCENTRATION AND PREDOMINANTLY SMALL LDL PARTICLES DESPITE SIMILAR TRADITIONAL LIPOPROTEIN LEVELS

ACC Poster Contributions

Ernest N. Morial Convention Center, Hall F

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Session Title: Genetic Associations with Cardiovascular Phenotypes and Endophenotypes

Abstract Category: 48. Genetics and Clinical Outcomes

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Background: Studies have suggested that carriers of the 719Arg allele of KIF6 have an increased risk of cardiovascular disease as well as an improved response to statins as compared to non-carriers. The association of the KIF6 polymorphism and LDL subclass distribution has not been reported.

Methods: We studied the distribution of lipoproteins according to KIF6 polymorphism in 218 apparently healthy non diabetic individuals (64% men, mean age: 61±14 years). KIF6 Trp719Arg genotypes were determined using an allele-specific real-time polymerase chain reaction genotyping assay. LDL subclass distribution was assessed using gradient gel electrophoresis and Lp(a) by immunoassay.

Results: In our study, 64% (n=142) and 36% (n=76) participants were carriers and non-carriers of KIF6 719Arg, respectively. No significant differences in baseline traditional risk factors were noted between KIF6 groups. However, carriers of 719Arg had a significantly higher Lp(a) level as well as a higher prevalence of predominantly intermediate (AB) and small LDL particles (B).

Conclusions: In this cohort of asymptomatic non-diabetic individuals, carriers of 719Arg had a higher concentration of LP(a) and higher likelihood of having smaller and dense LDL particles. This is the first study to report a possible association of the KIF6 polymorphism with atherogenic small LDL and if substantiated, may help explain the association of the KIF6 polymorphism with CHD risk.

| Association between KIF6 719 Arg status and Risk Factors | | | |
|--|--|--|---------|
| | Carriers Kif6 719Arg allele (n=142) | Non-carriers KIF6 Arg allele (n=76) | P value |
| LDL-C (mg/dl) | 96 ± 35 | 99 ± 40 | 0.6 |
| HDL-C (mg/dl) | 54 ± 15 | 56 ± 15 | 0.24 |
| TG (mg/dl) | 137 ± 86 | 124 ± 77 | 0.24 |
| Apo B (mg/dl) | 83 ± 23 | 79 ± 23 | 0.20 |
| Lp(a) (mg/dl) | 35 ± 46 | 22 ± 25 | 0.02 |
| LDL subclasses | | | |
| Large pattern A | 50% | 72% | 0.01 |
| Intermediate pattern AB | 29% | 14% | |
| Small dense pattern B | 21% | 14% | |